

Progress report

Factors affecting morbidity and mortality after surgery for obstructive jaundice: a review of 373 patients

A retrospective study of 373 patients undergoing surgery for relief of bile duct obstruction identified three independent factors associated with increased postoperative morbidity and mortality. These three factors were: (i) an initial haematocrit of 30% or less; (ii) an initial plasma bilirubin of greater than 200 $\mu\text{mol/l}$; (iii) a malignant obstructing lesion. The presence of two or all three of these risk factors identified a group of patients with a one third postoperative mortality and it is in this group of patients that preliminary non-operative or definitive non-operative biliary drainage might prove most useful.

Surgery for obstructive jaundice continues to be associated with significant morbidity and mortality despite recent advances both in preoperative diagnosis and postoperative care.^{1 2} Several studies have identified preoperative factors which define groups of patients at high risk of postoperative mortality³⁻⁵ and it has been suggested that mortality in these high risk groups might be reduced by the use of the newly available techniques of external or internal drainage of the obstructed biliary tree.^{1 6-8} The most comprehensive study identified eight preoperative factors which were associated with increased postoperative morbidity and mortality.³ The authors, however, did not take into account the effect of the operative procedure performed, half of all the deaths occurring in the minority of patients undergoing the most extensive surgical procedures, nor did they assess any interrelation of the eight risk factors.

A retrospective study was undertaken to assess effect of preoperative and operative parameters on postoperative outcome. It was the aim of our study to identify those factors which were independently associated with an increased risk of postoperative morbidity and mortality and which may be useful in predicting patients likely to benefit from preoperative and non-operative biliary drainage.

Methods

PATIENTS

Three hundred and seventy three patients with clinical jaundice (plasma bilirubin of greater than 50 $\mu\text{mol/l}$, normal range 2-18 $\mu\text{mol/l}$) who underwent surgery for relief of bile duct obstruction during the period 1976-81 in the department of Clinical Surgery, Royal Infirmary, Edinburgh were identified and a full review of all clinical case records undertaken.

Table 1 *Operative procedures performed in 373 patients with obstructive jaundice*

<i>281 patients with benign obstruction</i>	
255 : cholecystectomy + exploration of common bile duct	– 214 supraduodenal exploration – 41 combined supraduodenal + transduodenal exploration
10 : exploration of common bile duct only	
12 : simple bypass	
4 : resection of stricture	
<i>92 patients with malignant obstruction</i>	
6 : Whipple's procedure	
2 : hepaticojejunostomy	
71 : cholecystenterostomy	
13 : choledchoenterostomy	

Of the 373 patients, 281 had a benign and 92 a malignant cause of obstruction. The operative procedures performed in these two groups of patients are listed in Table 1. Two hundred and seventy of the operations were performed by a consultant surgeon, 91 by a senior registrar and 12 by a registrar. All patients received parenteral mannitol and vitamin K₁ before surgery. A total of 186 patients received antibiotics preoperatively, this being determined by the patient's clinical state and the wishes of the consultant in charge.

Full clinical, laboratory, and operative data were collected for all patients. Any postoperative complication was noted as was the date and cause of death. For the purposes of this study postoperative mortality was defined as death within 30 days of operation or death during the same hospital admission. Data on the following postoperative complications were available; renal failure, gastrointestinal bleeding, wound infection, wound dehiscence, septicaemia, abdominal abscess, and postoperative pancreatitis. Renal failure was defined as a rising urea in the postoperative period despite adequate fluid replacement. Gastrointestinal bleeding was noted as a complication only where the total measureable blood loss exceeded 500 ml.

Table 2 *Risk factors studied in group of 373 patients with obstructive jaundice*

*Age	*white blood cell count (WBCC)
*Sex	*bilirubin‡
*Pancreatitis†	*alanine aminotransferase‡ (ALT)
*Diabetes	*alkaline phosphatase‡ (AP)
*Haematocrit	*infected bile§
*ESR	*diagnosis
Length of history	Pyrexia
Weight loss	Platelet count
Abdominal pain	Prothrombin time ratio
History of smoking	Albumin‡
History of chest problems	Urea‡

* One of the 12 risk factors studied in every patient.

† A history of previous pancreatitis before admission.

‡ Measured in plasma.

§ Bile swabs taken in every case and where a positive growth was obtained patients were considered to have infected bile.

|| Final diagnosis of benign or malignant obstruction.

In an initial group of 120 patients, the relationship of 22 clinical, laboratory, and operative parameters to postoperative morbidity and mortality was analysed. A group of 12 factors was identified for more extensive study in the total group of 373 patients. Table 2 lists all 22 factors analysed, the 12 extensively studied being listed first. All biochemical and haematological values were those from the initial blood sample taken at the time of admission and in every case before the institution of any form of treatment. A number of patients with low haematocrits received preoperative blood transfusion.

The statistical methods used in the analysis were: (i) multiple logistic regression to test the relationship of preoperative factors and postoperative

Table 3 Risk factors and their relationship to postoperative mortality when fitted alone

Risk factor	Patients (%)	Mortality (%)	Significance
Age			
60 or less	34	3.9	
>60	66	13.4	p<0.05
Sex			
Female	63	7.3	
Male	37	12.2	NS
Pancreatitis			
Yes	8	0.0	
No	92	9.9	NS
Diabetes			
No	93	7.2	
Yes	7	34.5	p<0.0005
Haematocrit (35–54%)*			
>30	82	2.3	
30 or less	18	40.9	p<0.0005
ESR			
50 or less	75	7.1	
>50	25	15.2	p<0.05
WBCC ($4-10 \times 10^9/l$)*			
$10 \times 10^9/l$ or less	72	7.4	
$>10 \times 10^9/l$	28	13.6	NS
Bilirubin (2–18 $\mu\text{mol/l}$)*			
200 or less	73	3.7	
>200	27	23.5	p<0.0005
ALT (20–40 U/l)*			
100 or less	68	6.3	
>100	32	15.0	p<0.01
AP (100–400 U/l)*			
100 or less	33	0.8	
>100	67	13.1	p<0.01
Infected bile			
Yes	40	8.7	
No	60	9.4	NS
Disease			
Benign	75	3.7	
Malignant	25	26.1	p<0.0005

* Normal ranges shown in brackets.

mortality, (ii) Chi-squared tests to analyse the relationship of postoperative complications and mortality and the relationship of preoperative factors and postoperative complications, and (iii) Fisher's exact test where numbers were small.

Results

There were 34 deaths within 30 days of operation giving a hospital mortality of 9.1%. The operative procedure performed had no effect on postoperative mortality, with only one death occurring in the group of patients undergoing major surgical resections for malignant disease. Neither the experience of the operating surgeon nor the prior administration of antibiotics had an effect on postoperative mortality.

A univariate analysis of the relationship of the 12 factors to postoperative mortality identified eight which were significantly related. All 12 factors, their frequency and the relationship to mortality are shown in Table 3. When adjustments were made for the interrelation of factors as in a multivariate analysis only three remained significant, an initial haematocrit of 30% or less ($p < 0.0005$), an initial plasma bilirubin of over $200 \mu\text{mol/l}$ ($p < 0.01$) and a diagnosis of malignant obstruction ($p < 0.05$). There was no effect on mortality of preoperative blood transfusion in

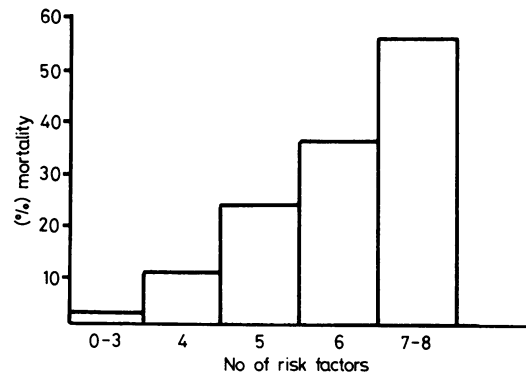


Fig. 1 Relationship of mortality and increasing numbers of unadjusted risk factors.

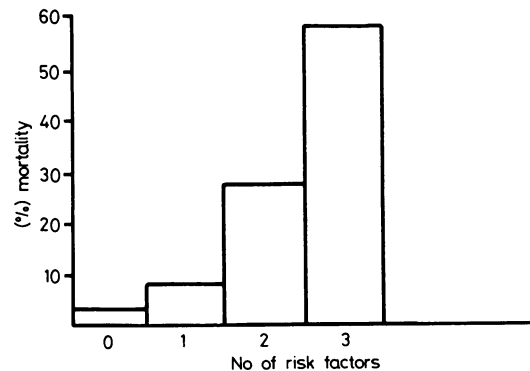


Fig. 2 Relationship of mortality and increasing numbers of adjusted factors.

patients with an initial low haematocrit. Exclusion of the diagnosis from the analysis did not lead to any other factor becoming independently significant, but the effect of bilirubin increased relatively in significance ($p < 0.0005$).

The correlation between the number of unadjusted risk factors per patient and mortality showed a significant increase in mortality when more than three factors were present, mortality increasing progressively in patients with increasing numbers of risk factors (Fig. 1). A similar stepwise progression was noted when mortality was compared with the three adjusted risk factors (Fig. 2). Increased mortality was seen in the 89 patients with two or three risk factors, with 29 out of the 34 deaths occurring in this group (Table 4), significantly different from the five deaths in the other 286 patients who had nil or one risk factor ($p < 0.0005$). One factor, a history of pancreatitis, was associated with a decreased likelihood of postoperative mortality ($p < 0.05$).

The frequency of the seven postoperative complications specifically studied and their relationship to postoperative mortality is shown in Table 5. Three complications were associated with an increased risk of mortality, these being renal failure ($p < 0.0005$), gastrointestinal bleeding ($p < 0.0005$), and abdominal abscess ($p < 0.05$). Thirteen patients developed renal failure, of whom eight died and 27 developed postoperative gastrointestinal haemorrhage with 13 deaths.

Analysis of the three adjusted risk factors and postoperative complications (Table 6) showed that the frequency of renal failure and gastrointestinal bleeding increased in direct proportion to the number of adjusted risk factors present. Four other complications, wound infection, wound dehiscence, septicaemia, and abdominal abscess were significantly more common in the group of patients with one or more risk factors. The

Table 4 Risk factors (adjusted) and their relationship of postoperative mortality

Risk factors (no.)	Patients (no.)	Deaths (no.)	Mortality (%)
0+1	286	5	1.7
2+3	87	29	33*

* Significantly different from groups of patients with 0 or 1 risk factors ($p < 0.0005$).

Table 5 Frequency of postoperative complications and their relationship to mortality

Complication	Frequency		Mortality	
	Patients (no.)	%	Patients (no.)	%
Renal failure	13	3.5	8	61.5†
GI bleeding	27	7.2	13	48.1†
Wound infection	36	9.7	3	8.3
Wound dehiscence	12	3.2	3	25.0
Septicaemia	24	6.4	6	25.0
Abdominal abscess	18	4.8	5	27.8*
Pancreatitis	14	3.8	2	14.3

Mortality significantly different between those with and those without the complication: * $p < 0.05$;

† $p < 0.0005$.

Table 6 Relationship of the three adjusted risk factors and postoperative morbidity

Risk factors (no.)	0	1	2	3
Patients (no.)	224	62	63	24
Complication	% patients presenting with this complication			
Renal failure	0.4	1.6	7.9‡	25.0§
GI bleeding	0.9	9.7‡	14.3§	41.7§
Wound infection	7.6	22.6‡	3.3	12.5
Wound dehiscence	0.9	6.5*	3.2	16.7§
Septicaemia	4.4	16.1‡	6.3	0.0
Abdominal abscess	0.4	19.3§	6.3‡	4.1
Pancreatitis	4.5	3.2	1.6	4.2

Significantly different from group of patients with 0 risk factors:

* $p < 0.05$; † $p < 0.01$; ‡ $p < 0.005$; § $p < 0.0005$.

three septic complications were most commonly seen in elderly patients with a benign obstructing lesion, a bilirubin of $>200 \mu\text{mol/l}$ and the presence of bacteria in the bile and thus it is not surprising that the frequency of these complications was highest in the group of patients with only one major risk factor. The only complication which was seen with equal frequency in all groups of patients was postoperative pancreatitis.

Discussion

Mortality rates of between eight and 33%^{1 2} have been reported for surgery to relieve bile duct obstruction. It has been postulated that decompression of the obstructed biliary tree by external or internal biliary drainage by reducing bilirubin concentrations will reduce postoperative mortality and morbidity.^{1 6} The procedures of internal or external non-operative drainage are not without complications^{1 6 9-11} and thus are most appropriate in patients at high risk of postoperative mortality.³ Attempts to identify such patients have studied a series of preoperative factors in isolation, but all have failed to adjust for the inter-relationship of these factors.³⁻⁵ In this present series three independent risk factors were identified which were associated with a significantly increased risk of postoperative mortality, these being an initial haematocrit of 30% or less, an initial plasma bilirubin of greater than $200 \mu\text{mol/l}$ and a diagnosis of malignant obstruction. Patients with two or all three of these risk factors represent a group at high risk with a one third postoperative mortality, and it is this group of patients in whom preoperative or non-operative drainage might prove most useful.

The most significant of the three factors was an initial low haematocrit and the influence of haematocrit was not reduced by preoperative blood transfusion. A low haematocrit is likely to signify some degree of malnutrition.³ A period of preoperative drainage without full attention to the nutritional status of the patient is thus unlikely to have any influence on the predictive value of the initial low haematocrit.⁶

Internal or external biliary drainage will reduce plasma bilirubin,^{1 6-8} which has been shown in this and other series to be an important factor in postoperative outcome.^{3 4} Reduction of plasma bilirubin below $200 \mu\text{mol/l}$ should on the basis of the present analysis lead to a reduction in operative mortality.

The third factor, the diagnosis of malignant obstruction cannot be influenced and thus the mortality of operating on patients with a malignant obstructing lesion will remain significantly higher than patients with a benign obstruction. Few patients are suitable for resection and in the majority palliation is all that can be performed.¹⁻⁴ In the latter group few patients survive longer than eight months.⁴ An alternative in this group of patients may be drainage with an internal biliary stent which appears to offer palliation similar to operative bypass, without the risk of operative mortality.⁷⁻⁸ Internal drainage, however, should only be performed after confirmation of the malignant nature of the obstruction by fine needle aspiration cytology.

Renal failure remains a problem in patients with obstructive jaundice,¹²⁻¹³ despite the use of mannitol. The frequency of this complication increases proportionately with the number of major risk factors present. The same association is seen with the other major complication related to mortality, gastrointestinal bleeding.⁴⁻¹⁴ Attempts to improve nutrition, lower plasma bilirubin and avoid an unnecessary operation in at least some patients with malignant obstruction, should theoretically reduce the incidence of these potentially lethal complications.

The studies on preoperative biliary drainage remain¹⁻⁶⁻¹⁰⁻¹⁵⁻¹⁶ at present inconclusive with little data on randomised prospective trials available.¹⁶ From this study three major risk factors for operative mortality have been identified. A period of preoperative drainage with an attempt to improve nutritional status and control infection for patients with a benign obstructing lesion, a haematocrit of 30% or less and a plasma bilirubin of over 200 $\mu\text{mol/l}$ should theoretically reduce operative mortality. Preoperative drainage can be achieved either by insertion of a percutaneous catheter, a duodenoscopically placed endoprosthesis or by endoscopic papillotomy. Similarly in patients with malignant disease and one or both of the other risk factors preoperative drainage (internal or external) or the avoidance of operation by the insertion of an internal stent with measures to improve nutrition and control infection should likewise reduce postoperative mortality. It remains to be seen whether more careful selection of patients for these drainage procedures will lead to a significant reduction in overall mortality and morbidity.

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References

- 1 Nakayama T, Ikeda A, Okuda K. Percutaneous transhepatic drainage of the biliary tract. Technique and results in 104 cases. *Gastroenterology* 1978; **74**: 554-9.
- 2 Buckwater JA, Lawton RL, Tidrick RT. Bypass operation for neoplastic biliary tract obstruction. *Am J Surg* 1965; **109**: 100-5.
- 3 Pitt HA, Cameron JL, Postier RG, Gadacz TR. Factors affecting mortality in biliary tract surgery. *Am J Surg* 1981; **141**: 66-72.
- 4 Sato T, Saitoh Y, Koyama K, Watanabe K. Preoperative determination of operability in carcinoma of the pancreas and the periampullary region. *Ann Surg* 1968; **168**: 876-86.
- 5 Braasch JW, Gray BN. Considerations that lower pancreaticoduodenectomy mortality. *Am J Surg* 1977; **133**: 480-4.
- 6 Denning DA, Molnar W, Carey LC. Preoperative percutaneous transhepatic biliary decompression lowers operative mortality in patients with obstructive jaundice. *Am J Surg* 1981; **141**: 61-5.
- 7 Osnes M, Odd G, Gronoeth H. Non operative internal drainage of obstructed common bile ducts. *Arch Surg* 1979; **114**: 862-5.
- 8 Ferrucci JT Jr, Mueller PR, Harbin WP. Percutaneous transhepatic drainage: technique, results and applications. *Radiology* 1980; **135**: 1-14.
- 9 Mori K, Misumi A, Suziyama M *et al*. Percutaneous transhepatic bile drainage. *Ann Surg* 1977; **185**: 111-5.
- 10 Burcharth F, Efsen F, Christiansen LA *et al*. Non-surgical internal biliary drainage by endoprosthesis. *Surg Gynecol Obstet* 1981; **153**: 857-60.
- 11 Armstrong CP, Taylor TV. Intrapleural leakage of bile complicating percutaneous transhepatic drainage of the obstructed biliary tree. *J R Coll Surg Edinb* 1982; **27**: 308-9.
- 12 Dawson JL. Postoperative renal function in obstructive jaundice: effect of mannitol diuresis. *Br J Surg* 1966; **53**: 979-85.
- 13 Allison MEM, Prentice CRM, Kennedy AC, Blumgart LH. Renal function and other factors in obstructive jaundice. *Br J Surg* 1979; **66**: 391-7.
- 14 Zollinger RM, Williams RD. Appraisal of progress in surgical therapy: surgical aspects of jaundice. *Surgery* 1956; **39**: 1016-30.
- 15 McPherson GAD, Benjamin IS, Habib HA, Bowley NB, Blumgart LH. Percutaneous transhepatic drainage in obstructive jaundice: advantages and problems. *Br J Surg* 1982; **69**: 261-4.
- 16 Hatfield ARW, Tobias R, Terblanche J *et al*. Pre-operative external biliary drainage in obstructive jaundice: a prospective controlled clinical trial. *Lancet* 1982; **2**: 896-9.